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A Consideration of the Social Dimensions and Implications of Neuroimaging Research in Global Health, as Related to the Theory-Laden and Theory-Generating Aspects of Technology

Keywords: Neuroimaging; neuroscience; social dimensions and implications; LMICs

Drawing on insights from sociology, anthropology, and the history of science and medicine, this paper considers some of the social dimensions and implications for neuroimaging research undertaken within low- and middle-income countries (LMICs). It highlights three key interconnected issues: (1) technologies for enhancing understandings of ill-health are theory-laden; (2) such technologies are theory-generating; and (3) studies of mental ill-health can also introduce new idioms for understanding subjective distress. The paper unpacks and explores these issues. It argues that the use of neuroimaging technologies in population research has the potential to contribute to solidifying – or even introducing – a biological (and specifically brain-based) understanding of mental ill-health within the communities under study. Examples from studies of neuroscience and society in various high-income countries (HICs) where neuroimaging is popular within public discourse illustrates how this can happen, and with what effects. The social dimensions and implications of neuroimaging are issues that all researchers using these technologies need to not only anticipate, but also explicitly plan for (and potentially seek to mitigate). Without adequate consideration, neuroimaging research carries with it particular risks in relation to extending the epistemological coloniality associated with HIC-sponsored studies conducted within LMIC settings.

Introduction

Within many high-income countries (HICs), research sponsors are actively seeking to cultivate not only technoscientific innovation within low- and middle-income countries (LMICs), but also the catalysis of social change. Such research can risk the reproduction of colonialism (Noxolo, 2017), and in particular epistemological colonialism (Hlabangane, 201; Ndlovu-Gatsheni, 2012). This paper explores some of the social dimensions and implications for neuroimaging research driven or supported by HICs and undertaken within LMICs. In particular, it draws on existing

sociological, anthropological, and historical research to highlight three key connected issues. First, technologies for enhancing understandings of ill-health are theory-laden. Second, such technologies are theory-generating. Third, studies of mental ill-health can also introduce new idioms for understanding subjective distress. Consideration of these issues (unpacked below) is vital when undertaking studies in settings wherein particular (psycho)pathological categories of research interest and the use of specific neuroimaging technologies to research them are marginal, nascent, or otherwise unfamiliar.

Exploring the Social Dimensions and Implications of Neuroimaging Research

Research generates not only new knowledge, but also new interactions between individuals and institutions - and potentially stimulates new legal and policy frameworks. Science is thus an intrinsically social enterprise, including in relation to how research agendas are set, teams built, and credit apportioned (Latour and Woolgar, 1986). Robust scholarship in sociology, anthropology, and history has extensively investigated the social dimensions of science. This is drawn upon below to explicate and illustrate some of the key social dimensions and implications of neuroimaging research.

Technology as theory-laden

Technologies are sponsored, developed, marketed, and operated by people within and across societies: technology, like science, is thus a social enterprise. Accordingly, it can carry with it the values and assumptions of the contexts in which it is imagined, incorporated, and implemented (Winner, 1980; Benjamin, 2019). This leads to the first key point advanced in this paper:

technologies for enhancing understandings of ill-health are theory-laden. By this, it is meant that biomedical technologies carry with them particular assumptions about the nature of the body and of pathology (Lock and Nguyen, 2011). These define the operations and potential utility of technologies.

In what senses are neuroimaging technologies like PET and fMRI theory-laden? For a start, they assume the relevance of the brain to the self and its pathologies (Dumit, 2004), with neuroimages becoming “sites for excavating biological reality/pathology” (Prasad, 2005: 292). As historians have demonstrated, the siting of subjectivity within the brain is a product of particular social and technological histories (Vidal, 2009). In the case of neuroscientific research into experiences

regarded in many HICs as psychiatric disorder, the use of technologies like fMRI almost inevitably come to further inflame debate within psychiatry and beyond around the place, role, and impact of (neuro)biology in aetiology (Bertorelli, 2016). In contexts where the brain is not, a priori, taken to be of singular import to constructions of selfhood, gaps can feasibly emerge between the perspectives of neuroscientists and those of collaborating clinicians and research participants. The questions such disjuncture pose include the extent to which participant autonomy could be compromised by this kind of mismatch between perceptions of selfhood, the brain, and the power and potential of neuroimaging (Pickersgill, 2011).

Technology as theory-generating

As well as being theory-laden, **technologies are theory-generating**. In other words, the use of technologies of visualisation help to embed ideas about the body and pathology within not only the research studies in which they are employed, but also within the understandings and perspectives of those who come into contact them. This might be directly - e.g., as study participants - or indirectly, such as through the media or through healthcare. Buchbinder (2015) provides a compelling example of how neuroimaging research is reimagined within US pain clinics when professionals communicate with adolescent patients. This includes through hopeful metaphors to young people that, for instance, “held the promise of possible futures in which new neural connections could erase the work of intractable pain” (Buchbinder, 2015: 310). In Australia, Barnett et al (2018; 2020) have demonstrated how the use of neuroscientific technologies within research into drug addiction has contributed to a discourse of a ‘hijacked brain’ within clinical addiction practice, which has helped to generate new moral positions for people living with addiction. In societies where individuals are often imagined as self-contained units whose actions are governed predominantly through ‘personal responsibility’, allusions to a ‘hijacked brain’ can absolve people for behaviour regarded as inappropriate or illegal. This can have beneficial effects, while also orientating recovery towards the recovering brain per se (as opposed to more psychosocial formulations).

Cohn’s (2010) UK-sited research with patient-participants in mental health research using neuroimaging demonstrates the theory-generating (or at least theory-reifying) potential of technologies like fMRI. In interviews following a scan where participants were granted a souvenir of a neuroimage, people commonly treated this as a straightforward picture of their brain that could precisely localise their condition, “allowing [the brain] to be conceived as an

alien pathological entity” (Cohn, 2010: 75; see relatedly Rapp, 2011). In their work in Canada with people diagnosed with mood disorders, Buchman and colleagues (2013) demonstrate how neuroimaging research can be read as both potentially destigmatising and legitimizing of experience *and* as likely to reify particular diagnoses and to more tightly couple the relationship between pathology and identity. The findings of Buchman et al (2013) exemplify the ambivalent nature of the social implications of neuroimaging. More specifically: the theories that are generated through technology use and which circulate within professional, patient, and popular discourses have neither straightforwardly positive nor negative effects. Consequently, they require careful and context-specific disentangling and engagement.

New idioms of distress

Alongside being theory-laden and theory-generating, **the use of neuroimaging technologies in studies of mental ill-health can also introduce new idioms for understanding subjective distress.** This potential relates to the purposes of the study for which neuroimaging is being used, and the other tools being employed as part of the experimental infrastructure. For example, diagnostic categorisations from the US Diagnostic and Statistical Manual of Mental Disorders (DSM) can be introduced through research practices. Yet, this text is “embedded in a very specific [US] biomedical epistemology” (Behrouzan, 2016: 27), and its introduction sometimes generates ambivalence and friction within clinical settings where “global psychiatry and local psychiatries” (Kitanaka, 2012: 81) are made to share space. New idioms for understanding subjective distress are themselves theory-laden: they contain remnants of the social and epistemic contexts of their generation. Like technologies, particular assumptions come with them about the nature of body and mind, and the ways these (inter)act to produce experiences of distress. Further, new idioms of distress can be theory-generating: they can be picked up by a range of people (i.e., not solely scientists and clinicians) and informally applied to others. Notions of distress crafted largely, though not exclusively, in North America can become accommodated elsewhere in the world in processes of what Duncan (2018) terms ‘psy-globalization’ (see relatedly Cox and Webb, 2015; Davar, 2014; Mills, 2013). In the UK, a key example of the adoption of US diagnostic terminology is talk of attention deficit hyperactivity disorder (ADHD) in educational contexts. Teachers sometimes informally describe students they consider to be particularly restless and impulsive as “a bit ADHD”; the pupils then come to be treated differently by teachers, with potentially detrimental effects (Hallett and Hallett, 2016: 15). The neurobiological understandings that underpin such informal characterisations are themselves

propelled through the use of neuroimaging technologies and the widespread cultural circulations of texts and iconography associated with these.

There is ample evidence of how DSM diagnostic categories like depression have become entwined with pre-existing categories (such as *shenjing shuairuo* in China; Lee, 1999) and psychopharmacuetical treatments interweaved with existing ontologies of (ill-)health (e.g., the reference of SSRIs as ‘mind food’ in parts of India; Ecks, 2014). As Kohrt et al (2016) observe, research is needed into how psychopharmaceuticals come to be assimilated into people’s lives following their introduction within contexts where they were previously less readily available. Particular attention must be paid to (a) how new patterns of consumption in LMICs relate to the introduction of categories of distress from HICs (e.g., the US), and (b) how neuroimaging technologies and the research and discourse they enable are purposively leveraged or implicitly presented as a means to make sense of these categories. This is especially important given concerns that attempts to innovate mental healthcare within LMICs can or might contribute to problematic forms of (bio)medicalization (Davar, 2014; Mills, 2018; Ventevogel, 2014; White and Sashidharan, 2014; Whitley, 2015). The extent to which neuroimaging research can act as a vehicle for introducing new idioms of distress, and contribute to forms of biomedical imperialism and colonialism (cf. Ugwu, 2019), should be subject to close consideration. However, such considerations cannot themselves become a vehicle for a form of epistemological coloniality within which it is simply assumed that concepts and technological from HICs will necessarily have transformative effects in LMICs.

Conclusion

Alongside the ethical and legal aspects and implications of neuroimaging technologies (Palk et al, 2020), the social dimensions and ramifications of research employing these tools also demands attention - whether studies make use of either long established or newer techniques like optical tomography (Fishell et al, 2020) and mobile MRI (Shen et al, in press). Through considerations of the existing social scientific evidence base, this paper has argued that the use of neuroimaging technologies can contribute to solidifying – or even introducing – a biological (and specifically brain-based) idiom for, and set of understandings of, mental ill-health and subjective distress.

As existing scholarship has demonstrated, such introductions and forms of reification have happened in a range of nations. At the same time, this research has also revealed that there are

limits to the extent to which accounts of the self and of distress have been neurobiologised as a consequence of the proliferation of neuroimaging (Barnett et al, 2020; Broer et al, 2020; Pickersgill et al, 2011). People - including, of course, neuroscientists themselves - are often subtle and creative in whether and how they ascribe significance to the brain and imaging technologies. However, caution is nevertheless warranted about disjunctures in understandings between scientists and study participants which could compromise the autonomy of the latter. Scientists also need to be mindful of how and in what ways so-called 'local' knowledge about the self is encouraged via imaging research to reform around purportedly 'global' neurobiological understandings. The salience of this issues relates to the wider potential for HIC-funded research in LMICs reproduce and extend forms of biomedical and epistemological colonialism (Kalinga, 2019; Ndlovu-Gatsheni, 2012).

This commentary resists offering straightforward guidelines for how researchers might deal with the proliferation of categories and ontologies beyond neuroimaging research sites. This is not least as a consequence of the heterogeneity of LMICs (and HICs), and hence the redundancy of advancing universal recommendations that are unlikely to always resonate sufficiently with the particularities of national contexts. Nevertheless, it is argued that these matters are part of the range of social and ethical issues that researchers need to not only anticipate but also explicitly plan for (and potentially seek to mitigate). Such consideration and planning will benefit from close collaboration with social scientists (Pickersgill et al, 2018) – as well, of course, with the communities with which neuroscientists should be co-producing their research questions and study designs (Nyirenda et al, 2020).

Scientists undertaking neuroimaging research need to be particularly mindful of the social dimensions and implications of their work. This is particularly the case when undertaking studies that: (a) employ imaging technologies in populations for whom the research techniques and the pathological categories of research interest are marginal, nascent, or otherwise unfamiliar (especially for groups where stigmatisation is a possible outcome of the movement of scientific findings into wider society; e.g. paediatric populations (Wedderburn et al, 2020; Xie et al, 2020) and older adults (Bachlie et al, 2020; Farina et al, 2020)); and (b) have the potential to induce uncertainties and even alarm among participants and their families (such as imaging research into infant neurodevelopment; Katus et al, 2020; Turesky et al, 2020).

References

Armstrong, D., Elliott, J., Hallett, F. and Hallett, G. (eds.) (2016) *Understanding Child and Adolescent Behaviour in the Classroom: Research and Practice for Teachers*, Cambridge: Cambridge University Press.

Bachli, M. B., Sedeño, L., Ochab, J. K., Piguet, O., Kumfor, F., Reyes, P., Torralva, T., Roca, M., Cardona, J. F. C., Gonzalez Campo, C., Herrera, E., Slachevsky, A., Matallana, D., Manes, F., García, A. M., Ibáñez, A., and Chialvo, D. R. (2020) 'Evaluating the reliability of neurocognitive biomarkers of neurodegenerative diseases across countries: a machine learning approach', *NeuroImage*, 208, 116456, <https://doi.org/10.1016/j.neuroimage.2019.116456>.

Barnett, A., Dilkes-Frayne, E., Savic, M., and Carter, A. (2018) 'When the brain leaves the scanner and enters the clinic: the role of neuroscientific discourses in producing the problem of "addiction"', *Contemporary Drug Problems*, 45, 3, 227-243.

Barnett, A., Pickersgill, M., Dilkes-Frayne, E., and Carter, A. (2020) 'Neural imaginaries at work: exploring Australian addiction treatment providers' selective representations of the brain in clinical practice', *Social Science & Medicine*, 255, 6, <https://doi.org/10.1016/j.socscimed.2020.112977>.

Bertorelli, T. E. (2016) 'Hope and doubt in the promise of neuroimaging: the case of autism spectrum disorder', *Health*, 20, 5, 505-522.

Behrouzan, O. (2016) *Prozak Diaries: Psychiatry and Generational Memory in Iran*, Stanford: Stanford University Press.

Benjamin, R. (2019) *Race After Technology: Abolitionist Tools for the New Jim Code*, Cambridge: Polity.

Broer, T., Pickersgill, M. and Cunningham-Burley, S. (2020) 'Neurobiological limits and the somatic significance of love: care-givers' engagements with neuroscience in Scottish parenting programmes', *History of the Human Sciences*, 33, 5, 85-109.

Buchbinder, M. (2015) 'Neural imaginaries and clinical epistemology: rhetorically mapping the adolescent brain in the clinical encounter', *Social Science & Medicine*, 143, 10, 304-310.

Buchman, D. Z., Borgelt, E. L., Whiteley, L., and Illes, J. (2013) 'Neurobiological narratives: experiences of mood disorder through the lens of neuroimaging', *Sociology of Health & Illness*, 35, 1, 66-81.

Cohn, S. (2010) 'Picturing the brain inside, revealing the illness outside: a comparison of the different meanings attributed to brain scans by scientists and patients', in: Edwards, J., Harvey, P., and Wade, P. (eds.) *Technologized Images, Technologized Bodies*, Oxford: Berghahn Books, pp. 65-84.

Cox, N. and Webb, L. (2015) 'Poles apart: does the export of mental health expertise from the Global North to the Global South represent a neutral relocation of knowledge and practice?', *Sociology of Health and Illness*, 37, 5, 683-697.

Davar, B. V. (2014) 'Globalizing psychiatry and the case of 'vanishing' alternatives in a neocolonial state', *Disability and the Global South*, 1, 2, 266-284.

Dumit, J. (2004) *Picturing Personhood: Brain Scans and Biomedical Identity*, Princeton: Princeton University Press.

Duncan, W. (2018) *Transforming Therapy: Mental Health Practice and Cultural Change in Mexico*, Nashville: Vanderbilt University Press.

Ecks, S. (2014) *Eating Drugs: Psychopharmaceutical Pluralism in India*, New York: New York University Press.

Farina, F. R., Emek-Savaş, D. D., Rueda-Delgado, L., Boyle, R., Kiiski, H. Yener, G., and Whelan, R. (2020) 'A comparison of resting state EEG and structural MRI for classifying Alzheimer's disease and mild cognitive impairment', *NeuroImage*, 215, 116795, <https://doi.org/10.1016/j.neuroimage.2020.116795>.

Fishell, A. K., Arbeláez, A. M., Valdés, C. P., Burns-Yocum, T. M., Sherafati, A., Richter, E. J., Torres, M., Eggebrecht, A. T., Smyser, C. D., and Culver, J. P. (2020) 'Portable, field-based neuroimaging using high-density diffuse optical tomography', *NeuroImage*, 215, 116541, <https://doi.org/10.1016/j.neuroimage.2020.116541>.

Hlabangane, N. (2017) 'On the coloniality of research in Africa: the case of HIV/AIDS in South Africa', in: Nhemachena, A. and Mawere, M. (eds.) *Africa at the Crossroads: Theorising Fundamentalisms in the 21st Century*, Bamenda: Langaa RPCIG, pp 61-106.

Jasanoff, S., Markel, G. E., Petersen, J. C., and Pinch, T. (eds.) (1995) *Handbook of Science and Technology Studies*, Thousand Oaks, CA: Sage Publications.

Kalinga, C. (2019) 'Caught between a rock and a hard place: navigating global research partnerships in the global South as an indigenous researcher', *Journal of African Cultural Studies*, 31, 3, 270-272.

Katus, L., Mason, L., Milosavljevic, B., McCann, S., Rozhko, M., Moore, S. E., Elwell, C. E., Lloyd-Fox, S., de Haan, M., Drammeh, S., Mbye, E., Touray, E., Ceesay, M., Jobarteh, B., Darboe, M. K., Austin, T., Prentice, A. (2020) 'ERP markers are associated with neurodevelopmental outcomes in 1–5 month old infants in rural Africa and the UK', *NeuroImage*, 210, 116591, <https://doi.org/10.1016/j.neuroimage.2020.116591>.

Kitanaka, J. (2012) *Depression in Japan: Psychiatric Cures for a Society in Distress*, Princeton: Princeton University Press.

Kohrt, B. A., Mendenhall, E., and Brown, P. J. (2016) 'How anthropological theory and methods can advance global mental health', *The Lancet Psychiatry*, 3, 5, 396-398.

Latour, B. and Woolgar, S. (1986) *Laboratory Life: The Construction of Scientific Facts*, 2nd edition, Princeton: Princeton University Press.

Lee, S. (1999) 'Diagnosis postponed: shenjing shuairuo and the transformation of psychiatry in post-mao China', *Culture, Medicine and Psychiatry*, 23, 3, 349-80.

Lock, M. and Nguyen, V-K. (2011) *An Anthropology of Biomedicine*, Chichester: Wiley-Blackwell.

Mills, C. (2018) 'From 'invisible problem' to global priority: the inclusion of mental health in the Sustainable Development Goals', *Development and Change*, 19, 3, 843-866.

Ndlovu-Gatsheni S J., 2012, *Coloniality of Power in Development Studies and the Impact of Global Imperial Designs on Africa*. *ARAS*, 33, 2: 48-73.

Noxolo, P. (2017) 'Decolonial theory in a time of the re-colonisation of UK research', *Transactions of the Institute of British Geographers*, 42, 3, 342-344.

Nyirenda, D., Sariola, S., Kingori, P., Squire, B., Bandawe, C., Parker, M., and Desmond, N. (2020) 'Structural coercion in the context of community engagement in global health research conducted in a low resource setting in Africa', *BMC Medical Ethics*, 21, 90, <https://doi.org/10.1186/s12910-020-00530-1>.

Palk, A., Illes, J., Thompson, P. M., and Stein, D. J. (2020) 'Ethical issues in global neuroimaging genetics collaborations', *NeuroImage*, 221, 117208, <https://doi.org/10.1016/j.neuroimage.2020.117208>.

Pickersgill, M. D. (2011) 'Research, engagement and public bioethics: promoting socially robust science', *Journal of Medical Ethics*, 37, 11, 698-701.

Pickersgill, M., Cunningham-Burley, S. and Martin, P. (2011) 'Constituting neurologic subjects: neuroscience, subjectivity and the mundane significance of the brain', *Subjectivity*, 4, 3, 346-365.

Pickersgill, M., Chan, S., Haddow, G., Laurie, G., Sridhar, D., and Cunningham-Burley, S. (2018) 'The social sciences, humanities, and health', *The Lancet*, 391, 10129, 1462-1463.

Prasad, A. (2005) 'Making images/making bodies: visibilizing and disciplining through Magnetic Resonance Imaging (MRI)', *Science, Technology, & Human Values*, 30, 2, 291-316.

Ventevogel, P. (2014) 'Integration of mental health into primary healthcare in low-income countries: avoiding medicalization', *International Review of Psychiatry*, 26, 6, 669-679.

Rapp, R. (2011) 'A child surrounds this brain: the future of neurological difference according to scientists, parents and diagnosed young adults', in: Pickersgill, M. and Van Keulen, I. (eds.) *Sociological Reflections on the Neurosciences*, Bingley: Emerald, pp. 3-26.

Ross G. White and Sashidharan, S. P. (2014) 'Towards a more nuanced global mental health', *The British Journal of Psychiatry*, 204, 6, 415-417.

Shen, F. X., Wolf, S. M., Bhavnani, S., Deoni, S., Elison, J., Fair, D. Geethanath, S., Garwood, M., Gee, M., Kay, K., Lim, K., Lockwood-Estrin, G., Luciana, M., Pelouquin, D., Rommelfanger, K., Schiess, N., Siddiqui, K., Torres, E., Vaughan, J. T. (in press) 'Emerging ethical issues raised by highly portable MRI research in remote and resource-limited international settings', *NeuroImage*.

Turesky, T., Xie, W., Kumar, S., Sliva, D. D., Gagoski, B., Vaughn, J., Zöllei, L., Haque, R., Kakon, S. H., Islam, N., Petri, W. A., Nelson, C. A., and Gaab, N. (2020) 'Relating anthropometric indicators to brain structure in 2-month-old Bangladeshi infants growing up in poverty: a pilot study', *NeuroImage*, 210, 116540, <https://doi.org/10.1016/j.neuroimage.2020.116540>.

Ugwu, C. (2019) 'Framing local attitudes to a modern health intervention in the neoliberal order – culturalism and malaria control in Southeastern Nigeria', *Journal of Asian and African Studies*, 54, 7, 1048-1065.

Vidal, F. (2009) 'Brainhood, anthropological figure of modernity', *History of the Human Sciences*, 22, 1, 5-36.

Wedderburn, C. J., Subramoney, S., Yeung, S., Fouche, J-P., Joshi, S. H., Narr, K. L., [Rehman](#) A. M., Roos, A., [Ipser](#) J., Robertson, F. C., Groenewold, N. A., Gibb, D. M., [Zar](#), H. J., Stein, D. J., and Donald, K. A. (2020) 'Neuroimaging young children and associations with neurocognitive development in a South African birth cohort study', *NeuroImage*, 219, 116846, <https://doi.org/10.1016/j.neuroimage.2020.116846>.

Whiteley, L., Borgelt, E., Stewart, S. E., and Illes, J. (2017) 'Parent perspectives on brain scans and genetic tests for OCD: talking of difficult presents, desired pasts, and imagined futures', *BioSocieties*, 12, 4, 471-493.

Winner, L. (1980) 'Do artifacts have politics?', *Daedalus*, 109, 1, 121-136.

Xie, W., Kumar, S., Kakon, S. H., Haque, R., Petri, W. A., and Nelson, C. A. (2020) 'Chronic inflammation is associated with neural responses to faces in Bangladeshi children', *NeuroImage*, 202, 116110, <https://doi.org/10.1016/j.neuroimage.2019.116110>.

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